Comparison of Esmolol to Nitroglycerine in Controlling Hypotension During Nasal Surgery

Nazal Cerrahide Kontrollü Hipotansiyon Sağlanmasında Esmolol ile Nitrogliserinin Karşılaştırılması

Ayla Guney, Fatma Nur Kaya, Belgin Yavascaoglu, Alp Gurbet, Nazan Has Selmi, Sener Kaya, Oya Kutlay Department of Anesthesiology and Reanimation, Faculty of Medicine, Uludag University, Bursa, Turkey

Abstract

Objective: The aim of this study was to compare esmolol to nitroglycerine in terms of effectiveness in controlling hypotension during nasal surgery.

Materials and Methods: After approval by our institutional Ethics Committee, 40 patients were recruited and randomized into two drug groups: esmolol (Group E) and nitroglycerine (Group N). In group E, a bolus dose of 500 µg/kg esmolol was administered over 30 sec followed by continuous administration at a dose of 25-300 µg/kg/min to maintain systolic arterial pressure at 80 mmHg. In group N, nitroglycerine was administered at a dose of 0.5-2 µg/kg/min.

Results: During the hypotensive period, systolic arterial pressure, diastolic arterial pressure, mean arterial pressure, and heart rate were decreased 24%, 33%, 27% and 35%, respectively, in group E (p<0.001, p<0.001, p<0.001, p<0.001) and were decreased 30%, 33%, 34% and 23%, respectively, in group N (p<0.001, p<0.001, p<0.001, p<0.001). The decrease in heart rate was higher in group E during the hypotensive period (p=0.048). During the recovery period, diastolic arterial pressure and heart rate were decreased 9% and 18%, respectively, in group E (p=0.044, p<0.001). Systolic arterial pressure, diastolic arterial pressure, and mean arterial pressure were decreased 7%, 3% and 7%, respectively, in group N (p=0.049, p=0.451, p=0.045).

Conclusion: Esmolol provides hemodynamic stability and good surgical field visibility and should be considered as an alternative to nitroglycerine.

Key Words: Anesthesia, Controlled hypotension, Esmolol hydrochloride, Nitroglycerin

Özet

Amaç: Çalışmamızın amacı nazal cerrahide esmolol ve nitrogliserinin kontrollü hipotansiyon oluşturmadaki etkinliklerinin karşılaştırılmasıdır.

Gereç ve Yöntem: Etik kurul onayı alındıktan sonra 40 olgu randomize olarak, esmolol (Grup E) veya nitrogliserin (Grup N) şeklinde iki gruba ayrıldı. Grup E'de esmolol 30 sn içinde 500 μg/kg bolus doz verilerek idamede 25-300 μg/kg/dk ve Grup N'de nitrogliserin 0.5-2 μg/kg/dk infüzyon hızında sistolik arter basıncı 80 mmHg olacak şekilde uygulandı.

Bulgular: Hipotansif dönemde sistolik arter basıncı, diastolik arter basıncı, ortalama arter basıncı ve kalp atım hızı değerlerinde Grup E'de %24, %33, %27 ve %35 (p<0.001, p<0.001, p<0.001) azalma izlendi. Hipotansif dönemde kalp atım hızı değerlerindeki azalma Grup E'de yüksekti (p=0.048). Derlenme döneminde Grup E'de diastolik arter basıncı ve kalp atım hızı değerlerinde %9 ve %18 azalma (p=0.044, p<0.001), Grup N'de ise sistolik arter basıncı, diastolik arter basıncı, ortalama arter basıncı değerlerinde %7, %3 ve %7 (p=0.049, p=0.451, p=0.045) azalma saptandı.

Sonuç: Esmololun kontrollü hipotansiyonda benzer hemodinamik değişiklik ve iyi cerrahi görüş alanı sağlaması nedeni ile nitrogliserine alternatif olarak kullanılabileceği kanısındayız.

Anahtar Kelimeler: Anestezi, Kontrollü hipotansiyon, Esmolol, Nitrogliserin

Introduction

Functional Endoscopic Sinus Surgery (FESS) is among the nasal surgical procedures that have increased in popularity since the late 1970s. However, serious complications such as hemorrhage, meningitis and damage to the optic nerve may occur due to the close proximity of blood vessels, nerves, and the orbital and intracranial cavities. The lack of complete

distinction between the anatomic structures associated with hemorrhaging in the surgical area during the procedure contributes to increased complications [1, 2].

Controlled hypotension (CH) at a moderate level (mean blood pressure - MAP>60 mmHg) is often preferred during nasal surgical procedures such as FESS and septorhinoplasty in order to reduce hemorrhaging and thus reduce complications via improved surgical field visibility [1, 3]

Received: December 30, 2011 / Accepted: April 09, 2012

Suitable patient positioning, positive pressure ventilation, medication, epidural or spinal anesthesia and high sympathetic block are among the methods used in the intentional reduction of blood pressure. The basic purpose is to reduce cardiac flow and/or systemic vascular resistance [4]. In addition to the commonly used sodium nitroprusside (SNP), nitroglycerine (NTG), and trimethaphan, CH can also be achieved through the administration of other agents such as prostaglandin E1, calcium canal antagonists, beta-blockers, fenoldopam, and angiotensin-converting-enzyme inhibitor (ACE) inhibitors [3].

Ease of administration, fast onset of efficiency, short half-life after discontinuation, rapid elimination of toxic metabolites, lack of a negative impact on vital organs, and expected and dose-dependent efficacy are the desired properties of the ideal hypotensive agent [3]. The use of vasodilators in CH has been reported to result in the possible development of tachyphylaxis and cyanide toxicity with SNP [3, 5]. In recent years, the rapid onset and short half-life of esmolol, as well as easy titration and close blood pressure control, has resulted in its use for CH [6-10].

There are no studies on the quality of the surgical field as they apply to nasal surgical procedures that compare the selective β -adrenergic receptor blocker esmolol and the vaso-dilator NTG. Therefore, our study compared the effects of CH as performed with esmolol and NTG in cases with planned FESS and septorhinoplasty for the purpose of evaluating surgical area quality, liver/kidney functions, and blood gas values.

Materials and Methods

Our study was conducted on 40 patients classified as American Society of Anesthesiologists (ASA) groups I and II who were scheduled for nasal surgery upon approval by our faculty's Ethics Committee and following receipt of express written consent from the patients. Non-cooperating patients with uncontrolled systemic diseases (DM, asthma and chronic obstructive lung disease), hepatic and renal failure, psychiatric diseases and disorders of the cardiovascular system, as well as those with a known drug allergy or substance addiction were excluded from the study.

All patients were monitored with electrocardiography (ECG). The following parameters were also monitored: non-invasive blood pressure, peripheral oxygen saturation (SpO₂) and end tidal CO₂ pressure (ETCO₂). A vascular line was prepared. First, a 0.9% NaCl infusion was started at a rate of 5 mL/kg/hr, and 0.03 mg/kg of midazolam was administered. A 20 G cannula was inserted into the radial artery on the non-dominant side for invasive blood pressure monitoring. All patients were administered 100% oxygen and 2.5 mg/kg/

IV propofol, 2 μg/kg of fentanyl and 0.6 mg/kg of rocuronium for induction. Normocapnic ventilation (ETCO, 32-35 mmHg) was performed after endotracheal intubation. In addition, 50% O₂-N₂O₂ 6% desflurane and IV remifentanil infusion was started at a rate of 0.1 µg/kg/min for anesthetic maintenance. Anesthetic depth was monitored by BIS (BIS Quatro™, Aspect Medical System, Newton MA, USA), and desflurane concentrations were adjusted to keep the BIS in the 40-60 range. Remifentanil infusion was administered upon titration at a 0.1-0.5 µg/kg/min dosage range. If the heart rate was 20% below the (HR) control values, the infusion dosage was decreased, and if the heart rate was 20% above the control values, it was increased. Muscular relaxation was obtained by intravenous administration of 0.1 mg/kg of rocuronium during the operation. The BIS values, MAC values and desflurane and remifentanil amounts administered were logged during anesthesia.

To provide increased convenience for the surgical team and to reduce the amount of bleeding, all patients were laid in an approx. 30° reverse Trendelenburg position and administered a standard dose of lidocaine-adrenaline (1: 100,000 adrenaline + 0.5% lidocaine, 1-2 mL) at the surgical site.

Forty patients were randomized by a sealed envelope method into two drug groups: esmolol (Group E, n=20) or NTG (Group N, n=20) administration prior to surgical incision.

An infusion for Group E was continued with a bolus dose of esmolol 500 $\mu g/kg/30$ sec and maintained at 25-300 $\mu g/kg/min$. For group N, a titration of NTG at 0.5-2 $\mu g/kg/min$ was administered. The blood pressure targets for both groups were systolic arterial pressure (SAP) 80 mmHg or mean arterial pressure (MAP) 60-65 mmHg, and infusion was discontinued at the end of the surgical procedure. The period was logged until the target values were achieved for all patients. The total esmolol and NTG doses administered throughout the surgical process were also recorded.

The initial SAP, diastolic arterial pressure (DAP), MAP, HR, SpO₂ and ETCO₂ of all patients were logged prior to drug infusion (control) in the pre-anesthetic period, during the hypotensive period, and during post-operative compilation.

To evaluate the effects of hypotensive agents on organ systems, aspartate transaminase (AST), alanine transaminase (ALT), lactate dehydrogenase (LDH), urea and creatinine values that indicate hepatic and renal functions were recorded during both the pre-operative and post-operative periods. In addition, pH, PO₂ and lactate values in the arterial blood gas were recorded prior to the beginning of hypotensive drug infusion, during the hypotensive period, and during post-operative compilation.

A 0.5-mg atropine IV was administered in the case of bradycardia (HR <40 bpm). The administered infusion dose in the case of MAP <60 mmHg was reduced by half, and infusion of

the hypotensive agent was discontinued when no response was obtained within 5 minutes. An ephedrine 10-mg IV was administered when required. Any side effects observed throughout the procedure were recorded.

The OP site was evaluated by the surgeon using the 0-5 point bleeding scale (0: no bleeding, 1: low bleeding-bleeding does not require aspiration, 2: low bleeding-bleeding requires intermittent aspiration, 3: low bleeding-bleeding requires frequent aspiration, 4: moderate bleeding-bleeding becomes serious when aspirator is withdrawn from the OP site, 5: serious bleeding-requires persistent aspiration, OP impossible) during the intra-operative period.

Side effects such as nausea/vomiting, hypotension/hypertension, bradycardia/tachycardia, gastrointestinal system disorders, blurry vision, allergy, headache and chest pain were considered during the post-operative period. Pre- and post-operative data were logged by an independent anesthetist not affiliated with the study.

All data were statistically analyzed using the SPSS 13.0 (SPSS Inc., Chicago IL.) statistical software package. When comparing cases administered esmolol or NTG, a significant difference between the groups concerning hypotension required 19 cases per group (power 0.84, α =0.05, β =0.16). Data normality was analyzed through the Shapiro-Wilk test. A t-test was applied to compare both groups when the data were normally distributed. Data not normally distributed were compared using a Mann-Whitney U test. Comparison of the dependent groups was performed with the matching t-test for data with a normal distribution and a Wilcoxon Signed Rank test for nonnormally distributed data. Categorical data were examined by Pearson chi-square and Fisher's exact chi-square tests. The level of significance difference was set at p<0.05.

Results

The demographic and surgical data of the patients in Groups E and N as well as their target SAP achievement times, BIS values, and consumption of desflurane and remifentanil were found to be similar (Table 1).

The initial SAP, DAP, MAP, HR, SpO₂ and ETCO₂ values of the patients during pre-anesthetic induction were similar (Table 2).

The comparison between the values during the pre-hypotensive period and hypotensive agent infusion period, in particular SAP, DAP, MAP, and HR, revealed decreases of 24%, 33%, 27% and 35%, respectively (p<0.001, p<0.001, p<0.001 and p<0.001, respectively) for Group E, and 30%, 33%, 34% and 23%, respectively (p<0.001, p<0.001, p<0.001 and p<0.001, respectively) for Group N. Compared to the hypotensive agent infusion period, a significant reduction of 9% and 18% (p=0.044 and p<0.001, respectively) was observed during compilation of the DAP and HR control periods, respectively, of Group E, while the reduction

Table 1. Demographic data of patients, intraoperative BIS values, and drug consumption

	Group E (n=20)	Group N (n=20)	р
Gender (F/M)	12/8	13/7	0.807
Age (year)	29.80±7.80	32.30±10.70	0.738
Weight (kg)	69.10±12.30	72.20±12.60	0.436
Height (cm)	169.40±8.40	172.00±9.10	0.353
ASA (I/II)	19/1	17/3	0.605
Operation time (min)	92.70±21.20	106.00±25.50	0.705
Anesthesia time (min)	109.70±19.50	120.20±24.30	0.080
Target MAP time (sec)	68.00±8.30	75.00±11.70	0.143
BIS value	48.20±4.90	45.90±4.70	0.350
Desflurane consumption (MAC)	0.70±0.30	0.70±0.40	1.000
Remifentanil (µg/kg/min)	0.14±0.01	0.13±0.02	0.052

Data are given as n or mean±SD

MAP: Mean arterial pressure, MAC: Minimum alveolar concentration, BIS: Bispectral index score

Table 2. Baseline values before anesthesia induction

	Group E (n=20)	Group N (n=20)	р
SAP (mm Hg)	111.5±20.5	115.6±20.8	0.538
DAP (mm Hg)	71.9±12.6	72.9±12.1	0.799
MAB (mm Hg)	83.6±14.9	86.6±14.3	0.519
HR (bpm)	84.2±13	83.1±14.3	0.800
SpO ₂ (%)	99.1±0.8	99.3±0.9	0.462
ETCO ₂ (mm Hg)	34.2±2.6	35.2±3.7	0.328

Data are given as n or mean±SD

SAP: Systolic arterial pressure, DAP: Diastolic arterial pressure, MAP: Mean arterial pressure, HR: Heart rate, SpO₂: Peripheral oxygen saturation, ETCO₂: End-tidal carbon dioxide

in SAP, DAP and MAP values of Group N were 7%, 3% and 7%, respectively (p=0.049, p=0.451 and p=0.045, respectively). A comparison between average SAP, DAP and MAP values and the values prior to hypotensive infusion yielded no difference between the groups. The decreases in the HR values during the hypotensive period were found to be high (p=0.048) in Group E. No significant difference was observed in the group and intergroup comparisons of the SpO₂ and ETCO₂ values (Table 3).

Evaluation of surgical site hemorrhaging during the hypotensive period did not yield any significant difference between the groups.

Table 3. Before infusion (control), hypotensive period and recovery period data

	Group E (n=20)	Group N (n=20)	р
SAP (mmHg)			
Before infusion	116.7±17.8	129.6±25.5	0.713
Hypotensive period	86.5±7.9**	86.7±4.3**	0.921
Recovery period	117.1±11.0	117.2±18.2*	0.983
DAP (mmHg)			
Before infusion	74.2±12.4	80.3±12.7	0.132
Hypotensive period	50.5±9.0**	51.0±5.3*	0.733
Recovery period	72.2±6.4*	71.9±14.0*	0.931
MAP (mmHg)			
Before infusion	88.8±14.2	96.4±16.3	0.124
Hypotensive period	61.4±8.4*	61.7±5.2**	0.892
Recovery period	85.7±8.8	87.1±14.7*	0.716
HR (atım dk⁻¹)			
Before infusion	93.6±14.4	86.1±10.6	0.068
Hypotensive period	59.7±7.2**	64.7±8.3**	0.048
Recovery period	74.6±11.1*	79.2±13.3	0.242
SpO ₂ (%)			
Before infusion	99.3±0.7	99.4±0.6	0.630
Hypotensive period	99.2±0.6	99.3±0.5	0.570
Recovery period	99.3±0.5	99.3±0.6	1.000
ETCO ₂ (mm Hg)			
Before infusion	31.6±3.3	32.2±2.9	0.545
Hypotensive period	31.1±3.3	31.4±2.9	0.761
Recovery period	34.7±6.2	36.4±7.1	0.424
		· · · · · · · · · · · · · · · · · · ·	

Data are given as n or mean±SD

SAP: Systolic arterial pressure, DAP: Diastolic arterial pressure, MAP: Mean arterial pressure, HR: Heart rate, SpO₂: Peripheral oxygen saturation, ETCO₃: End-tidal carbon dioxide

*p<0.05 and **p<0.001, intra-group comparison, compared to before infusion

The comparison between the hepatic and kidney function parameters of the patients did not yield any statistically significant differences in the ALT, AST and urea values during the pre- and post-operative periods. The increase in the post-operative creatinine and LDH values compared to the pre-operative control values was found to be similar in both groups (p=0.003, p=0.024 for Group E and p=0.003, p=0.005 for Group N) (Table 4).

No significant difference in arterial blood samples received during the hypotensive period and compilation period in terms of pH, PO₂ and lactate values was found

Table 4. Preoperative and postoperative hepatic and renal functions

	Group E (n=20)	Group N (n=20)	n
ALT (/ .ll)	(11—20)	(11—20)	р
ALT (mg/dL)			
Preoperative	20.0±12.0	24.1±32.2	0.596
Postoperative (Day 1)	17.7±6.8	22±21.7	0.403
AST (mg/dL)			
Preoperative	21.1±9.0	21.8±12.5	0.840
Postoperative (Day 1)	21.8±9.5	23.0±12.3	0.731
Urea (mg/dL)			
Preoperative	24.6±9.4	25.0±7.5	0.882
Postoperative (Day 1)	27.0±8.7	24.4±5.5	0.265
Creatinine (mg/dL)			
Preoperative	0.7±0.2	0.7±0.1	1.000
Postoperative (Day 1)	0.9±0.2*	0.9±0.2*	1.000
LDH (mg/dL)			
Preoperative	122.4±16.7	124.5±14.5	0.673
Postoperative (Day 1)	136.3±20.6*	138.5±15.7*	0.706
Data are given as n or mean±SD AST: Aspartate transaminase, ALT: Alanine transaminase, LDH: Lactate dehydrogenase			

*p<0.05, intra-group comparison, compared to preoperative values

between the groups prior to hypotensive drug infusion. While the group assessments yielded a statistically significant decrease in the PO_2 during the pre-hypotensive drug infusion period, (p=0.039 for Group E and p<0.001, p=0.032, p<0.001 for Group N), the increase in the lactate values during the hypotensive and compilation period was also found to be statistically significant for both groups compared to the pre-drug infusion period (p=0.032, p=0.043 for Group E and p=0.027, p=0.017 for Group N) (Table 5).

A category ASA I patient in Group E developed bradycardia following esmolol bolus dose administration, and 0.5 mg atropine was administered intravenously. Esmolol infusion was initiated after stabilization of the hemodynamic parameters, and no further pre-operative complications were observed.

Except for nausea/vomiting, no post-operative side effects were observed. Nausea/vomiting was observed in 6 patients in Group E (30%) and 3 in Group N (15%).

Discussion

Our study compared the effects of the short-acting β -adrenergic receptor blocker esmolol and the vasodilator NTG for HR control under general anesthesia (desflurane and

Table 5. Arterial pH, PaO, and lactate values

	Group E (n=20)	Group N (n=20)	р
рН			
Before infusion	7.40±0.05	7.40±0.30	1.000
Hypotensive period	7.40±0.04	7.30±0.40	0.272
Recovery period	7.30±0.04	7.30±0.50	1.000
PaO ₂ (mmHg)			
Before infusion	273.70±65.07	278.20±68.20	0.832
Hypotensive period	247.40±57.60°	238.90±80.20*	0.702
Recovery period	152.30±67.50°	147.40±36.10*	0.776
Lactate (mmol/L)			
Before infusion	9.00±3.60	10.10±5.70	0.470
Hypotensive period	16.00±7.65*	16.00±7.60*	1.000
Recovery period	14.70±6.01*	20.10±10.40*	0.051
Data are given as the mean+SD			

Data are given as the mean±SD

remifentanil) in patients with a planned nasal surgical procedure. Anesthetic depth in both groups was maintained at a stable rate with BIS monitoring.

A close relationship between reduced MAP value and surgical site quality in surgical procedures has been shown [11]. A 38% reduction in hemorrhaging in 6298 patients who underwent FESS with HR control has been reported [12]. Cincikas and Ivaskevicius [13] have shown reduced bleeding and improved surgical view quality with 50 to 60 mmHg MAP maintenance an endoscopic nasal surgery with NTG (0.79 \pm 0.34 μ g/kg/min).

In their study comparing the effects of esmolol (500 µg/ kg load, 100-300 μg/kg/min) and SNP (0.25-4.0 μg/kg/min) on HR control (target MAP 55-65 mmHg) during orthognathic surgery, Blau et al. [14] reported the average MAP as 58.7±0.7 mmHg with esmolol and 61.8±0.4 mmHg with SNP. Furthermore, they suggested that 40% of the values observed in the esmolol and 53% in the SNP group remained outside of the target MAP range and that the reduction of blood loss with esmolol was more effective and controlled. Boezaart et al. [15] reported that while the surgical conditions during FESS with moderate hypotension (MAP>65 mmHg) and SNP (0.25 µg/kg/min) were not good, optimum surgical area was achieved with esmolol (500 µg/kg load, 100-300 µg/kg/min). The authors suggested that increased hemorrhage in the mucous membrane associated with the vasodilator effect of SNP and increased cardiac flow accounted for this result. The esmolol-related improvements in surgical conditions, however, were associated with the effective vasoconstriction of precapillary sphincters and arterioles in the mucous membranes.

Pilli et al. [16] reported that the use of esmolol at an average infusion rate of $330\pm10~\mu g/kg/min$ in tympanic surgery provided a bloodless surgical site with a 28.7% decrease in SAP, 26.5% in MAP and 33.4% in DAP. In addition to the reductions of a similar nature in blood pressure induced by esmolol and NTG in our study, a similar improvement was also observed in surgical site quality. We believe that the lower dose of the medication administered compared to other studies resulted from a minimized stress response to surgical stimulants, along with the reverse Trendelenburg positioning achieved through BIS monitoring.

Yoshikawa et al. [17] reported that the target MAP 60-70 mmHg during an NTG-accompanied mandibular osteotomy was achieved at 1-5 μ g/kg/min by an infusion rate of 13.3 \pm 1.5 min. Degoute et al. [8] reported that the target SAP (80 mmHg) in tympanoplasty procedures was achieved in 53.3 \pm 4.4 seconds with esmolol (210 \pm 33 μ g/kg/min). Using esmolol at 50-300 μ g/kg/min, Turan et al. [9] demonstrated that MAP (>60 mmHg) was achieved in 13.35 \pm 8.4 min. Although in our study, the period in which the target SAP (80 mmHg) achieved was statistically similar, it was achieved in a shorter period of time in the esmolol group (Group E: 68.0 \pm 8.3 sec, Group N: 75.0 \pm 11.7 sec).

SNP and NTG are frequently preferred for HR control during procedures. Yet, SNP can cause vasodilatation, reflex tachycardia and increased cardiac flow. When compared to SNP, the negative effects of NTG on reflex tachycardia, "rebound" hypertension, and organ perfusion - along with risk of overdose - are reported to be lower [18].

In contrast, esmolol causes a dose-dependent, controlled and slow decrease in blood pressure, with no observable tachycardia, unlike the deep fluctuations in blood pressure observed with SNP administration. In a study comparing esmolol and SNP in the achievement of target HR in patients scheduled for lumbar fusion and cerebrovascular surgery, Ornstein et al. [19] reported a 15.9% CAD increase in the SNP and 12.1% decrease in esmolol groups. They also suggested that discontinuation of the hypotensive agents resulted in a significant increase in MAP in the SNP group. Increases in CAD at 30 and 60 min were reported upon NTG application in a different study [17]. When comparing CAD control values, the decrease in the hypotensive and compilation period was 35% and 18% in the esmolol and 23% in the NTG groups. Our study did not find a significant decrease in the compilation period. In addition, bradycardia responsive to treatment after bolus administration in the esmolol group was observed in only one subject. Although a decrease in blood pressure after discontinuation of the drug infusions was lower in the NTG group, no difference was found between the groups.

 $[\]ensuremath{^{\circ}}\xspace p {<} 0.05,$ intra-group comparison, compared to before infusion values

The mechanisms that keep microcirculation under control should be taken into consideration while implementing techniques to reduce blood flow and provide a dry operation site. Auto-regulation through minimal tissue metabolism assumes the role of a local protective mechanism against excess reduction in blood flow. HR control should target sustainable protective auto-regulatory mechanisms, while deep hypotension should be avoided. Deep hypotension reducing organ perfusion may be associated with morbidity and mortality (0.02-0.06%) [3]. Therefore, our study refrained from deep HR control (MAP = 50 mmHg), and the target MAP was maintained in the 60-65 mmHg range. In a study where they researched the impact of NTG on the liver during HR control (MAP approx. 60 mmHg), Fukusaki et al. [20] found increased AST and LDH activity even though the patients were maintained in the normal range. Kol et al. [10] found that esmolol did not affect liver and kidney functions during HR control where the target MAP was 65-75 mmHg. Piper et al. [7] reported that the alpha-glutathione-D-transferase (Alpha GST) level in the blood samples collected from patients with and without the administration of esmolol and SNP, and deep hypertension (MAP 50-55 mmHg) in the endonasal sinus surgery, was higher in the hypotensive group and that such difference would return to normal by the second hour of the procedure. However, standard liver enzymes (alanine aminotransferase, aspartate aminotransferase, gamma-glutamyl transferase) were unchanged. Therefore, they suggested that deep hypotension could result in hepatocellular damage at a minimal level.

In their study researching the effects of isoflurane, esmolol and NTG on splanchnic perfusion during a hypotensive procedure (30% of control SAP, MAP>50 mmHg) of elective maxillofacial surgery, Andel et al. [21] suggested that gastric intra-mucous pH remained unchanged, no increase in the blood lactate levels was observed, and thus, organ perfusion was sustained. In an approx. 2-hour HR control procedure where SAP was performed at 70 to 79 mmHg, Nagata et al. [22] reported that the urinary NAG (n-acetylglucosaminidase) activity indicative of tubular renal function and gamma-GTP activity were higher at 3 post-operative days, and reduced post-op creatinine clearance, free water clearance and urinary volume returned to normal within 3 days. It was suggested that mildly reversible renal tubular cell damage could be observed. However, serum SGOT and SGPT levels were shown to remain unchanged throughout the 21-day post-operative period. Our moderate HR control procedure demonstrated that while liver functions were not affected by standard tests, the increase in post-operative creatinine and lactate values in both groups was similar but within normal ranges. However, we believe that the possible minimal effects on organ functions require more elaborate testing in cases of risk.

Degoute et al. [8] reported that the changes in PaO₂, PCO₂ and lactate in the arterial blood gas samples in the administration of remifentanil, SNP and esmolol for HR control (SAP 80 mmHg) in patients with planned tympanoplasty were similar. Pilli et al. [16] reported that there was no significant difference in the arterial blood gas samples in hypotensive anesthesia achieved with esmolol (28.7% reduction in SAP). In their study comparing NTG and SNP with normo-tensive anesthetic procedures in mandibular osteotomies, Yoshikawa et al. [17] demonstrated a similar reduction in the PaO_a values in the blood gas samples taken from of all three groups 60 minutes post-surgery. They suggested that the reductions in the PaO₂ value were not affected by the administration of both drugs and that this could be explained by the ventilation/perfusion mismatch associated with artificial ventilation during anesthetic administration. Moreover, they considered the lack of difference in increased lactate associated with tissue hypoxia and the fact that pH, BE and HCO, values remained within the normal range to be proof of tissue oxygenation continuing throughout hypotension. In addition to similar hemodynamic changes, our study yielded no significant difference in arterial blood gas and lactate values between the groups.

Consequently, we believe that esmolol can be safely used as an alternative to NTG. Its HR control-related surgical site quality in nasal surgical procedures is superior, and its effects on hepatic/renal functions and arterial blood gas parameters are similar to NTG. In addition, esmolol produces easy control of blood pressure with no "rebound" tachycardia and hypertension.

Conflict of interest statement: The authors declare that they have no conflict of interest to the publication of this article.

References

- Baker AR, Baker AB. Anaesthesia for endoscopic sinus surgery. Acta Anaesthesiol Scand 2010; 54: 795-803.
- 2. Ulualp SO. Complications of endoscopic sinus surgery: appropriate management of complications. Curr Opin Otolaryngol Head Neck Surg 2008; 16: 252-9.
- 3. Degoute CS. Controlled hypotension: a guide to drug choice. Drugs 2007; 67: 1053-76.
- 4. Kayhan Z. Klinik Anestezi. Genişletilmiş 3. Baskı. Ankara: Logos Yayıncılık; 2004, 496-501.
- 5. Dietrich GV, Heesen M, Boldt J, Hempelmann G. Platelet function and adrenoceptors during and after induced hypotension using nitroprusside. Anesthesiology 1996; 85: 1334-40.
- Gökçe BM, Karabiyik L, Karadenizli Y. Hypotensive anesthesia with esmolol. Assessment of hemodynamics, consumption of anesthetic drugs, and recovery. Saudi Med J 2009; 30: 771-7.
- Piper SN, Haisch G, Kumle B, et al. Effects of esmolol- and sodium nitroprusside-induced controlled hypotension on hepatocellular integrity in patients undergoing endonasal sinus surgery].

- Anasthesiol Intensivmed Notfallmed Schmerzther 2003; 38: 781-6
- Degoute CS, Ray MJ, Manchon M, Dubreuil C, Banssillon V. Remifentanil and controlled hypotension; comparison with nitroprusside or esmolol during tympanoplasty. Can J Anaesth 2001; 48: 20-7.
- Turan G, Dincer E, Ozgultekin A, et al. Comparison of dexmedetomidine, remifentanil and esmolol in controlled hypotensive anaesthesi. The Internet Journal of Anesthesiology 2008; 17: 2.
- Kaygusuz K, Yildirim A, Kol IO, Gursoy S, Mimaroglu C. Hypotensive anaesthesia with remifentanil combined with desflurane or isoflurane in tympanoplasty or endoscopic sinus surgery: a randomised, controlled trial. J Laryngol Otol 2008; 122: 691-5.
- Sieskiewicz A, Drozdowski A, Rogowski M. The assessment of correlation between mean arterial pressure and intraoperative bleeding during endoscopic sinus surgery in patients with low heart rate. Otolaryngol Pol 2010; 64: 225-8.
- 12. Maune S, Jeckström W, Thomsen H, Rudert H. Indication, incidence and management of blood transfusion during sinus surgery: a review over 12 years. Rhinology 1997; 35: 2-5.
- Cincikas D, Ivaskevicius J. Application of controlled arterial hypotension in endoscopic rhinosurgery. Medicina 2003; 39: 852-9.
- 14. Blau WS, Kafer ER, Anderson JA. Esmolol is more effective than sodium nitroprusside in reducing blood loss during orthognathic surgery. Anesth Analg 1992; 75: 172-8.

- Boezaart AP, van der Merwe J, Coetzee A. Comparison of sodium nitroprusside- and esmolol-induced controlled hypotension for functional endoscopic sinus surgery. Can J Anaesth 1995; 42: 373-6.
- Pilli G, Güzeldemir ME, Bayhan N. Esmolol for hypotensive anesthesia in middle ear surgery. Acta Anesthesiol Belg 1996; 47: 85-91.
- Yoshikawa F, Kohase H, Umino M, Fukayama H. Blood loss and endocrine responses in hypotensive anaesthesia with sodium nitroprusside and nitroglycerin for mandibular osteotomy. Int J Oral Maxillofac Surg 2009; 38: 1159-64.
- 18. Endrich B, Franke N, Peter K, Messmer K. Induced hypotension: Action of sodium nitroprussid and nitroglyserin on the microcirculation. Anesthesiology 1987; 66: 605-13.
- Ornstein E, Matteo RS, Weinstein JA, Schwartz AE. A controlled trial esmolol for induction of delibrate hypotension. J Clin Anesth 1988; 1:31-5
- 20. Fukusaki M, Matsumato M, Yamaguci K. Effect of hemodilution during controlled hypotension on hepatic, renal and pancreatic function in humans. Clin Anesth 1996; 8: 545-50.
- 21. Andel D, Andel H, Hörauf K, Felfernig D, Millesi W, Zimpfer M. The influence of deliberate hypotension on splanchnic perfusion balance with use of either isoflurane or esmolol and nitroglycerin. Anesth Analg 2001; 93: 1116-20.
- 22. Nagata N, Agune T, Takeshita M, Yoshikawa G, Takasaki M. Effects of induced hypotension on perioperative renal and hepatic functions. Masui 1993; 42: 1157-61.